

**REVERSAL OF PROINFLAMMATORY RESPONSE BY
LIGATING THE MACROPHAGE Fc γ RI RECEPTOR**

Abstract of the Disclosure

5 Ligation of the Fc γ receptor type I (Fc γ RI) on IL-10-producing cells
leads to a selective upregulation of IL-10 production, which in turn induces a marked
suppression of IL-12 biosynthesis by IL-12-producing cells, particularly macrophages.
The ligation of the Fc γ RI receptor thus downmodulates IL-12 production via a
mechanism that is dependent on macrophage-derived IL-10. Agents for ligating Fc γ RI
comprise, for example, multivalent antibodies which bind the Fc γ RI receptor, immune
10 complexes comprising antibodies which contain the Fc region of IgG, and IgG
multimers, preferably IgG dimers and trimers. The ligating agent may be administered
to therapeutically inhibit proinflammatory immune responses. In particular, the ligating
agent may be administered to treat or prevent endotoxic shock associated with bacterial
endotoxemia, and to treating autoimmune disorders.